Fig. 1A

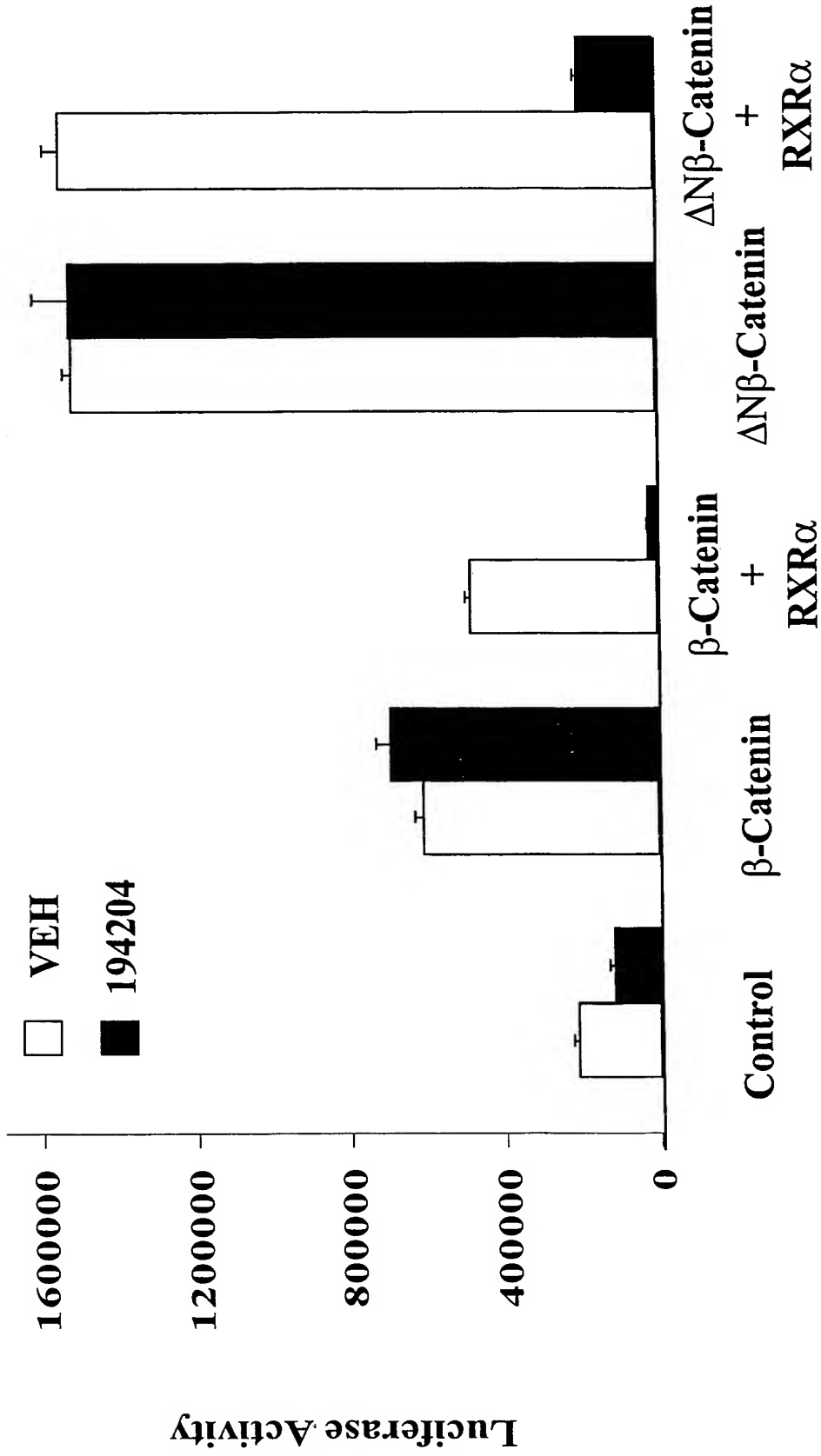

AGN194204 Inactivates β-catenin-mediated Gene Transactivation *via* RXRα
VEH
194204
Luciferase Activity
1600000
1200000
800000
400000
0
Control
β-Catenin
β-Catenin + RXRα
ΔNβ-Catenin + ΔNβ-Catenin + RXRα

Fig. 1B

Reduction of Stable β-catenin Transactivation by Stable Expression of RXRα in 293 Cells

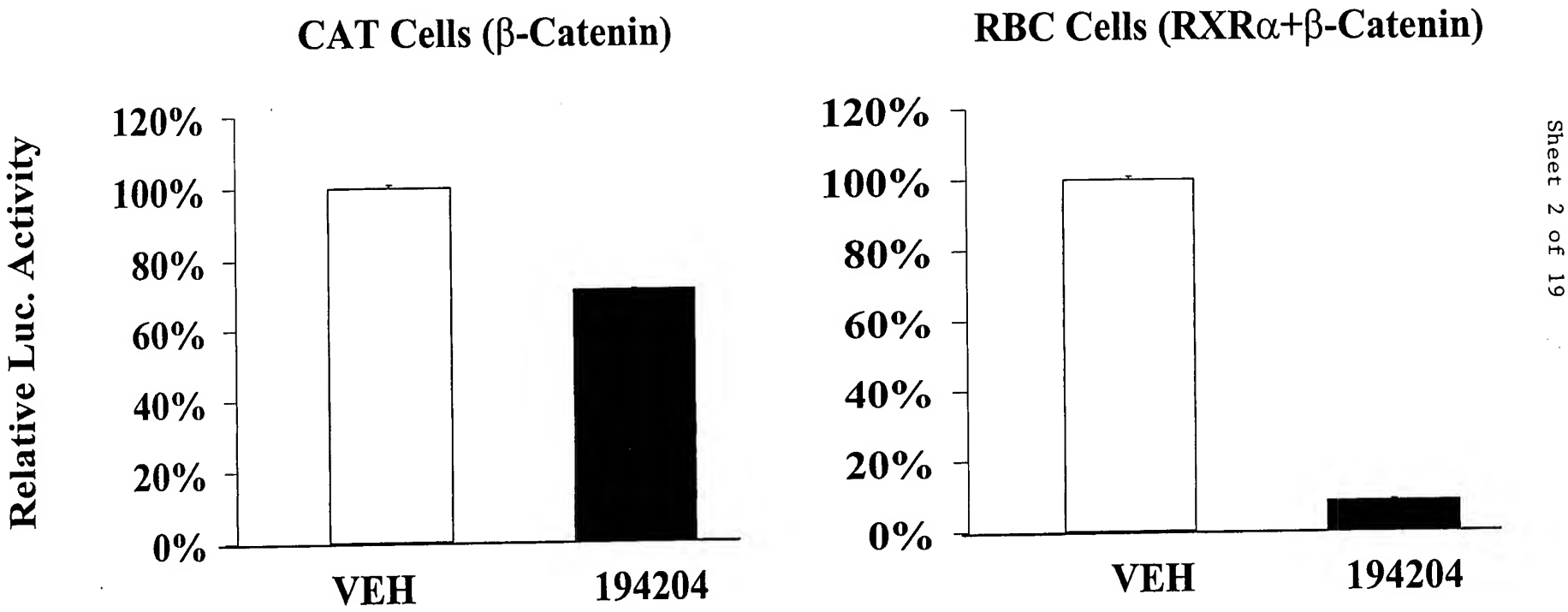


Fig. 1C

Inactivation of Endogenous β-Catenin in Gene Transactivation by AGN194204 *via* RXRα in Colon Cancer Cells

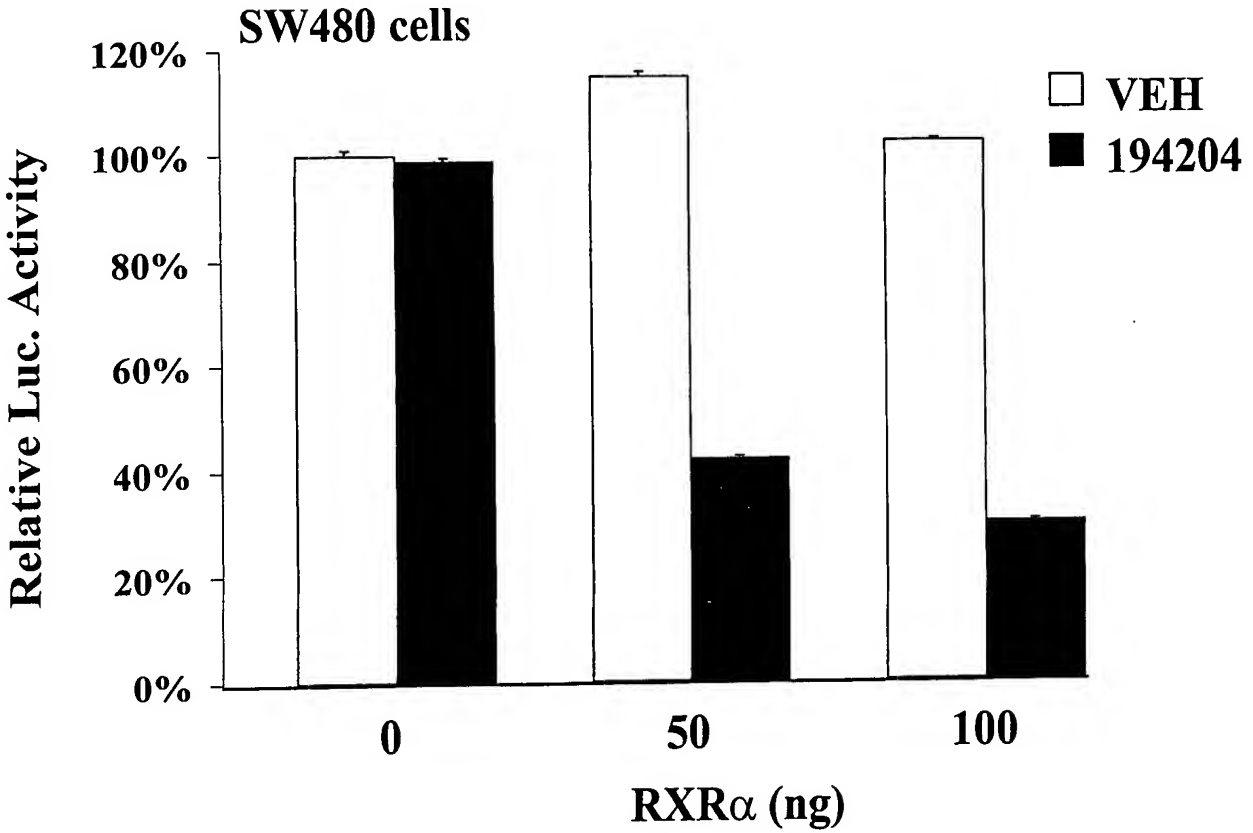


Fig. 2A

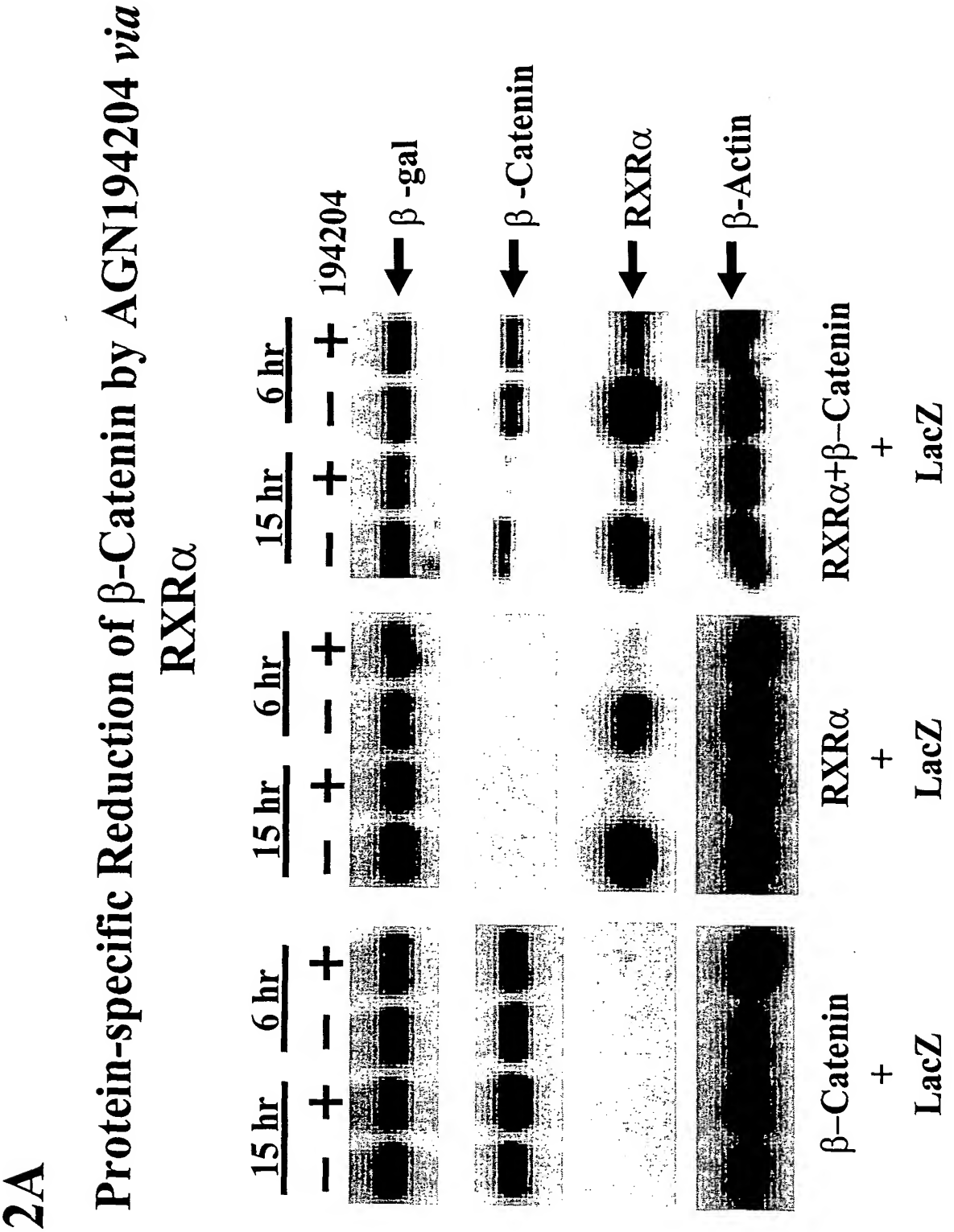

Protein-specific Reduction of β-Catenin by AGN194204 via RXRα
15 hr
6 hr
15 hr
6 hr
194204
β -gal
β -Catenin
RXRα
β-Actin
β–Catenin + LacZ
RXRα + LacZ
RXRα+β–Catenin + LacZ

Fig. 2B

Time Course of Degradation of β-Catenin & RXRα by AGN194204 in Stable Cell Line RBC

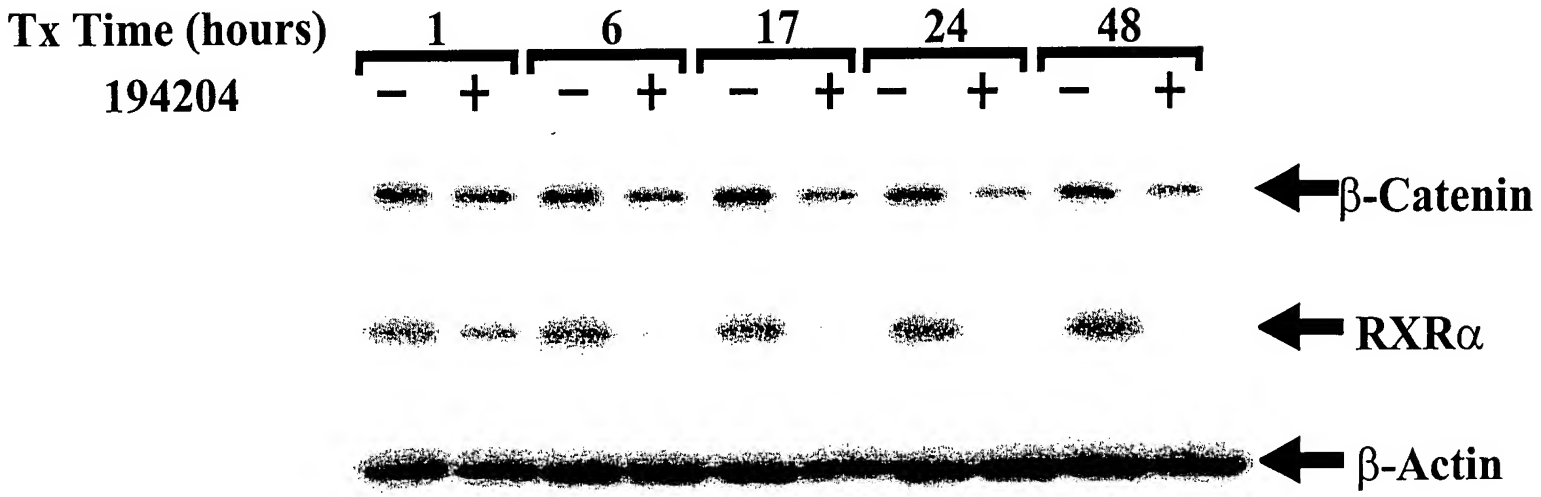


Fig. 2C

RXRα Is Stoichiometrically Required for Reduction of β-Catenin

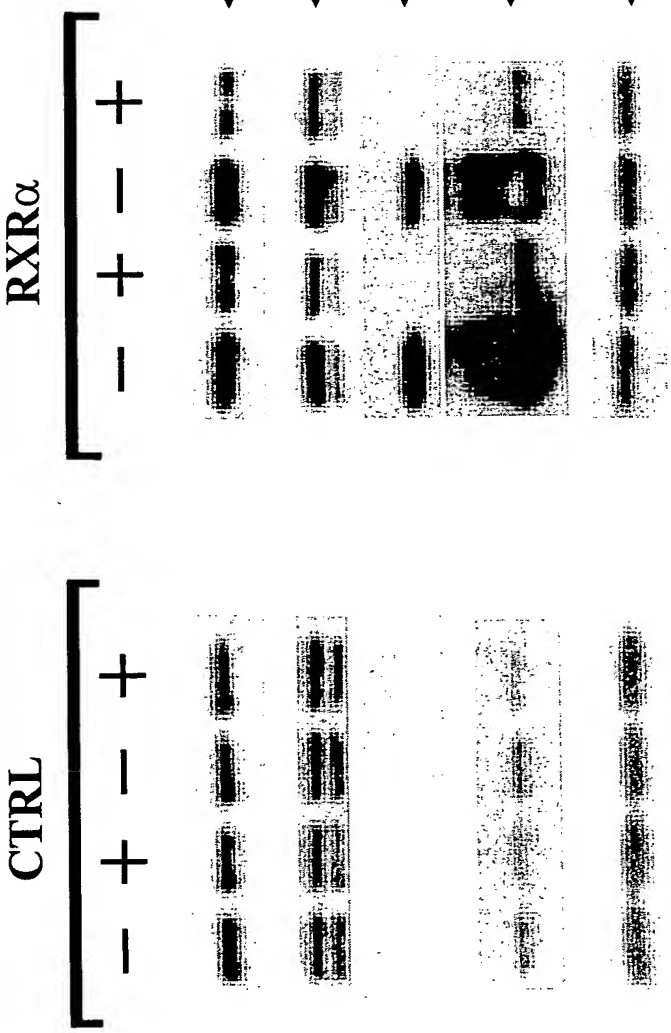


Fig 2D

RXRα-dependent Reduction of β-Catenin by AGN194204 - Stoichiometric Requirement for RXRα

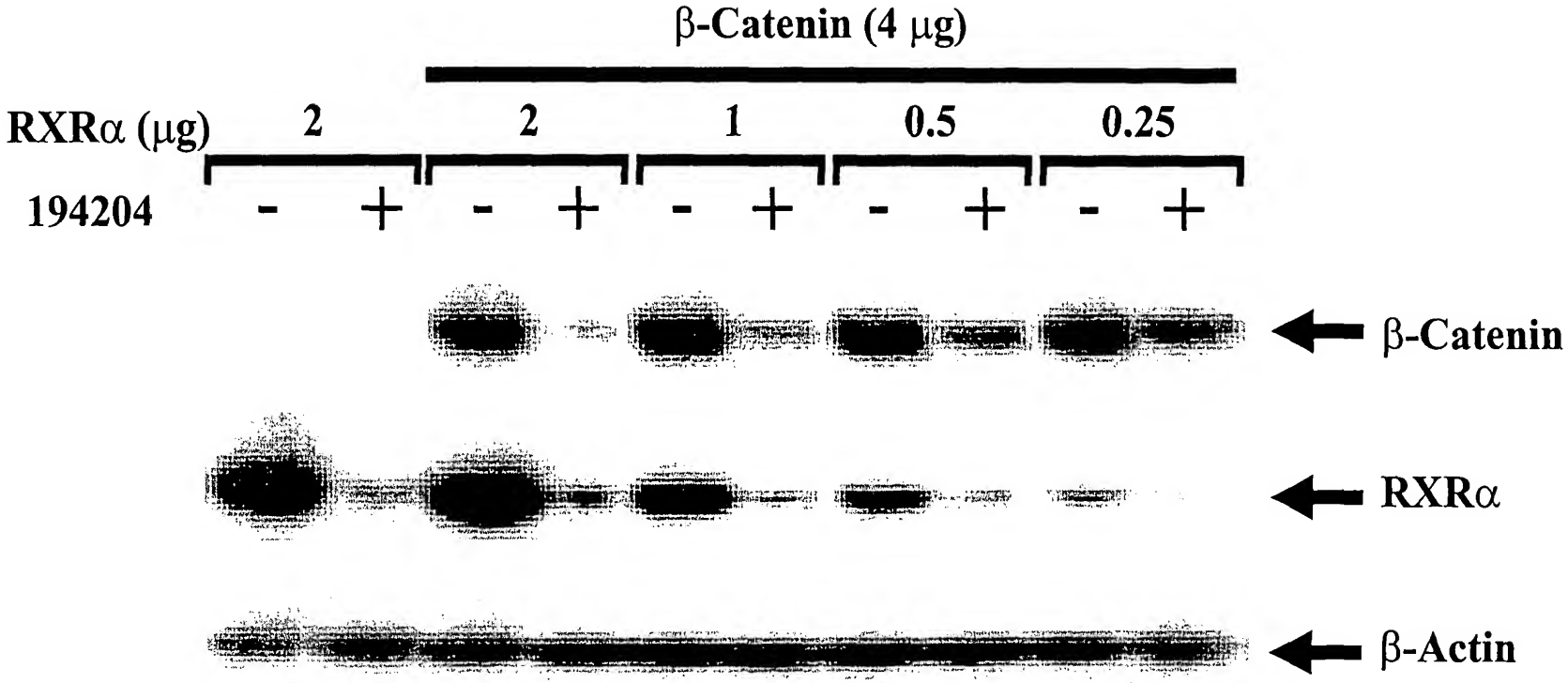


Fig. 2E

AGN194204 Ubiquitously Reduces Wild Type & Mutant β-Catenins

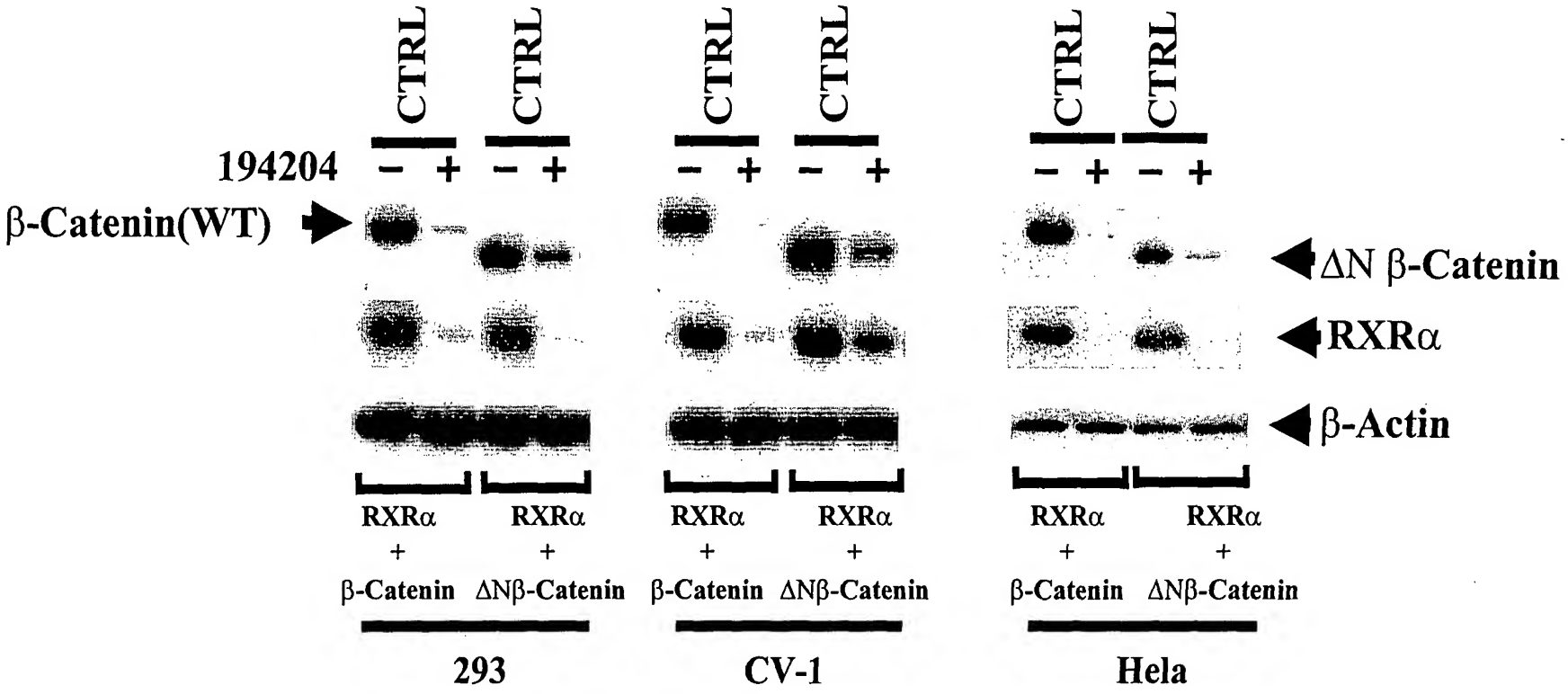


Fig. 2F

AGN194204 Does Not Affect the β-Catenin mRNA Level in Stable Cell Line RBC

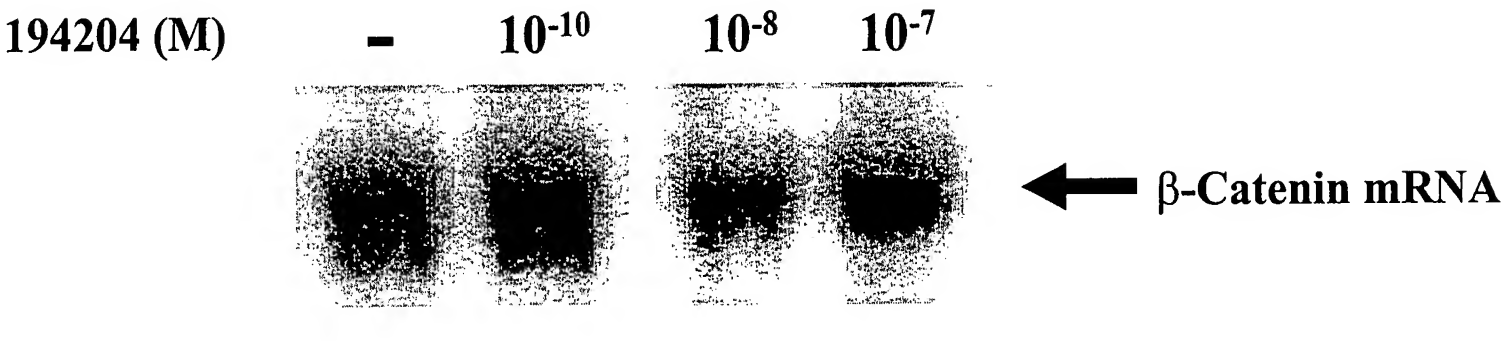


Fig. 3

Time Course of Degradation of ΔNβ-Catenin by AGN194204 in Stable Cell Line RmBC

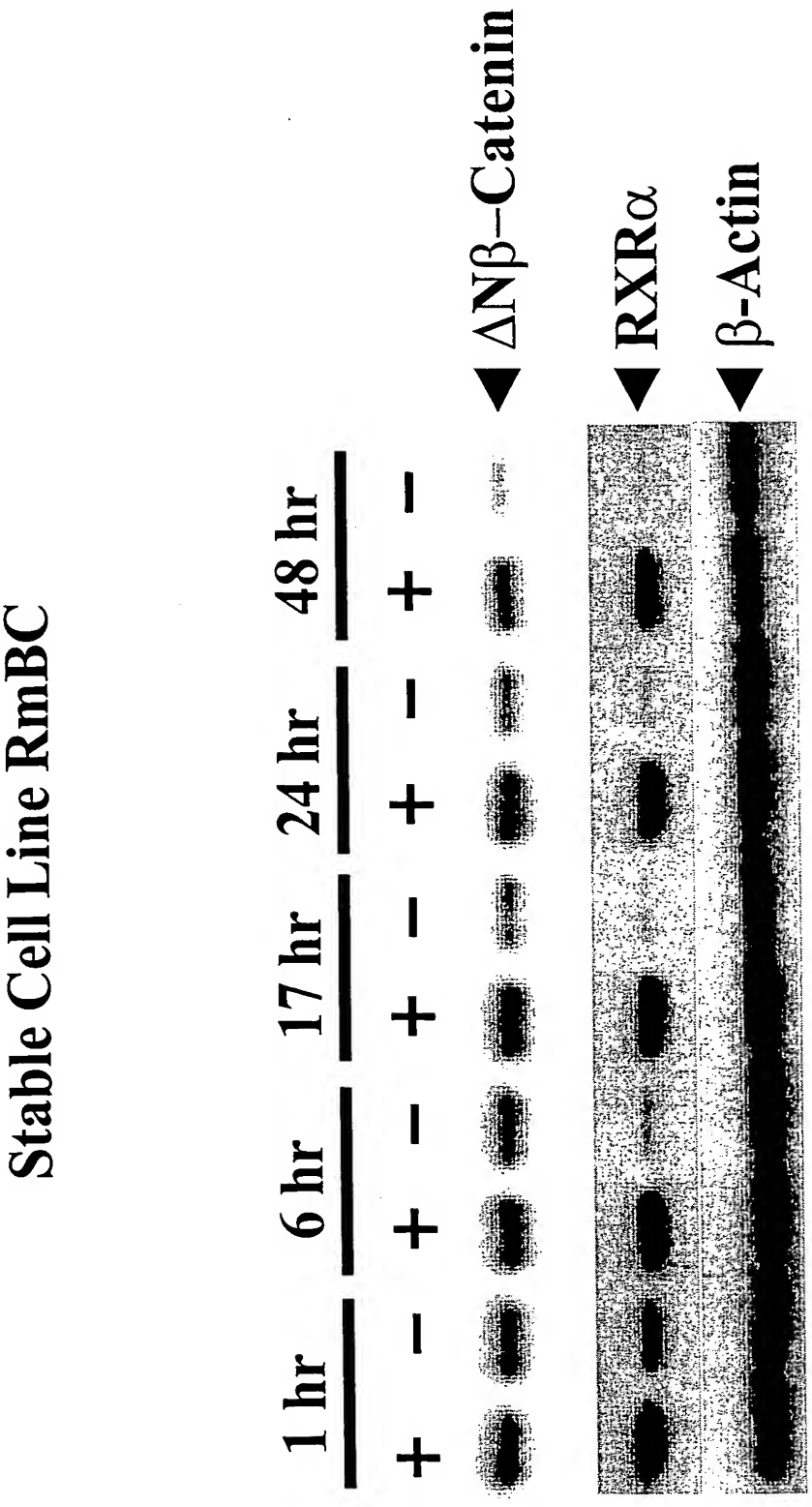


Fig. 4A

High Potency of AGN194204 in Reduction of RXRα and β-Catenin Proteins & Its Antagonism by AGN195393

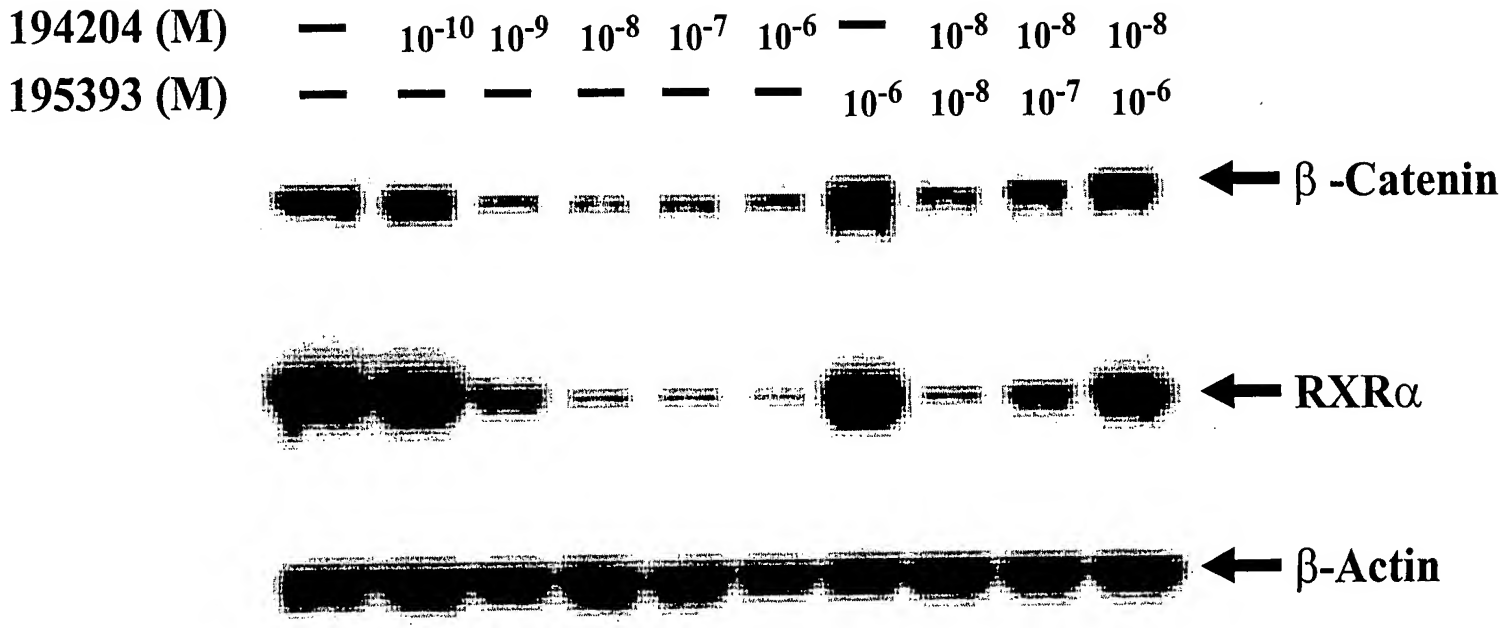


Fig. 4B

Ligand-specificity in Reduction of RXRα and β-Catenin Proteins

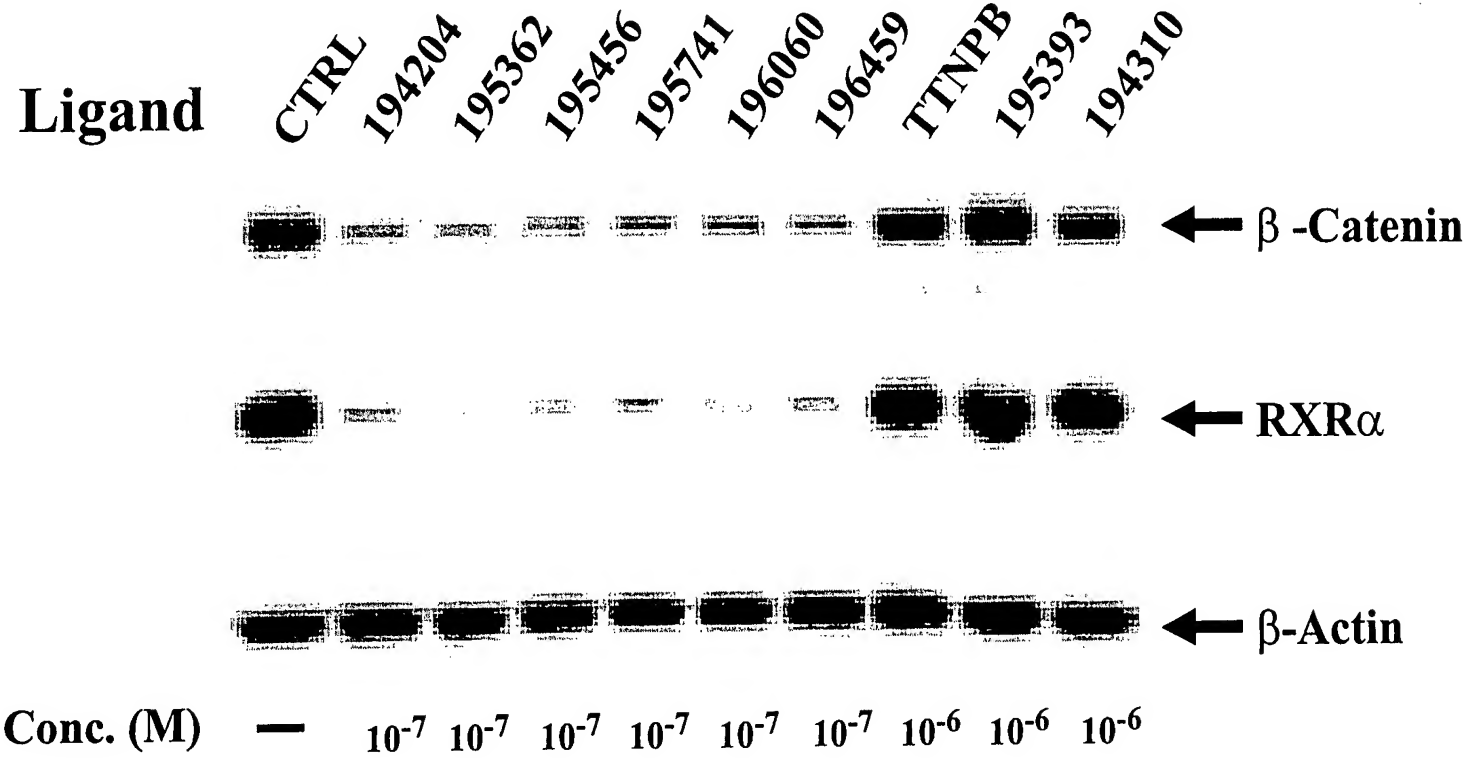


Fig. 4C

AGN194204-induced Reduction of RXRγ & β-Catenin Proteins

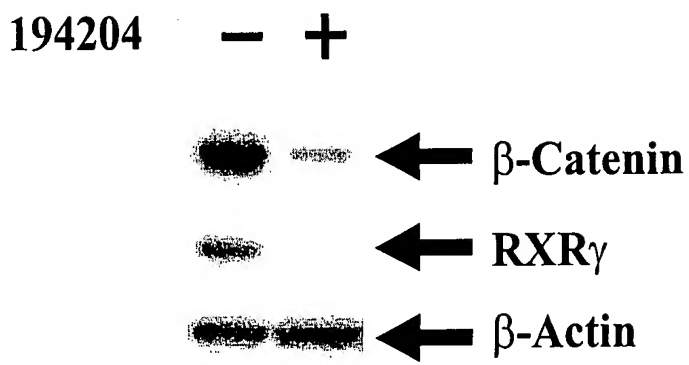


Fig. 4D

RARα but not β and γ Minimally Reduces β-Catenin Protein

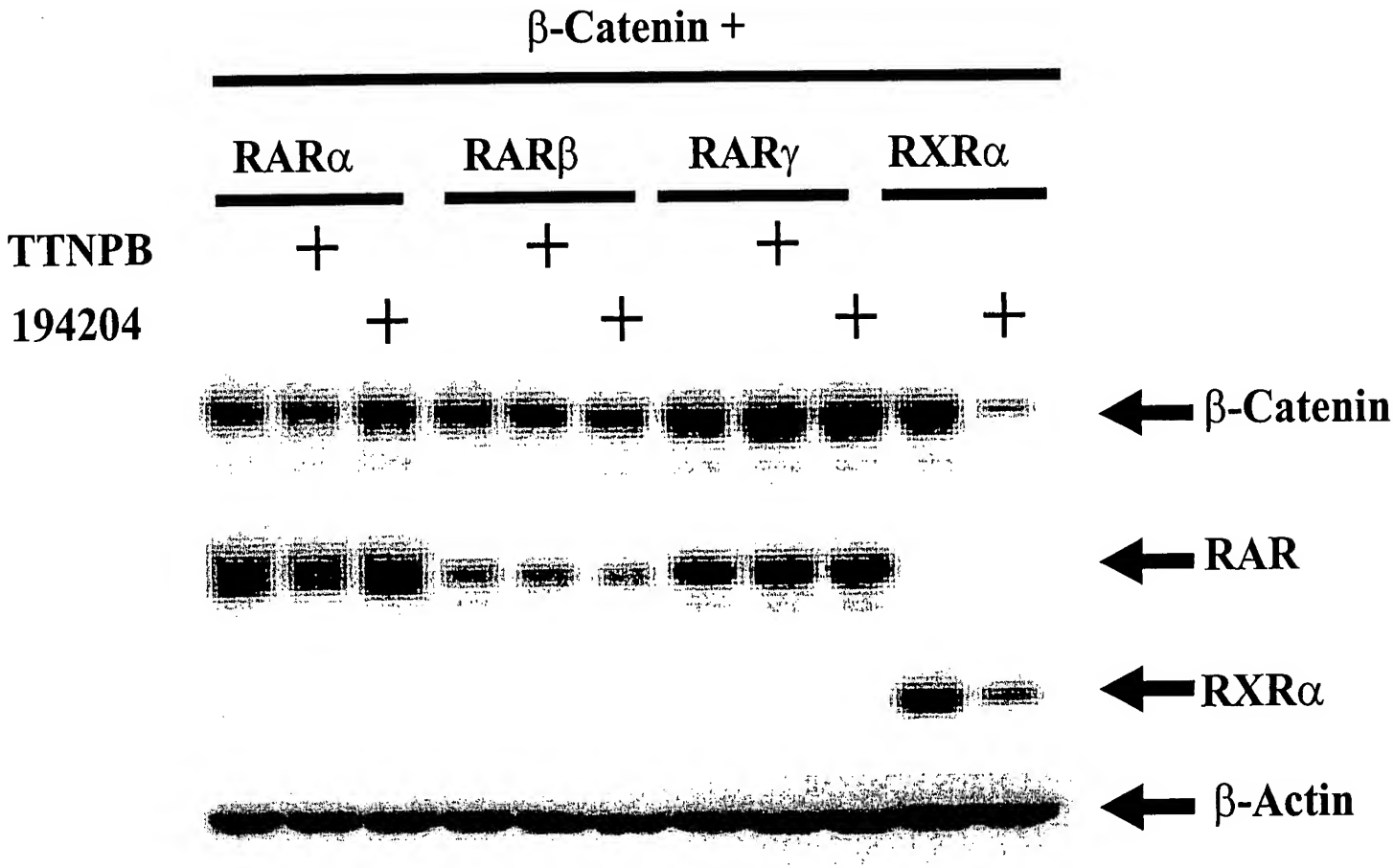


Fig. 4E

RXRα & Its Ligand Reduce Both RAR and β-Catenin Proteins

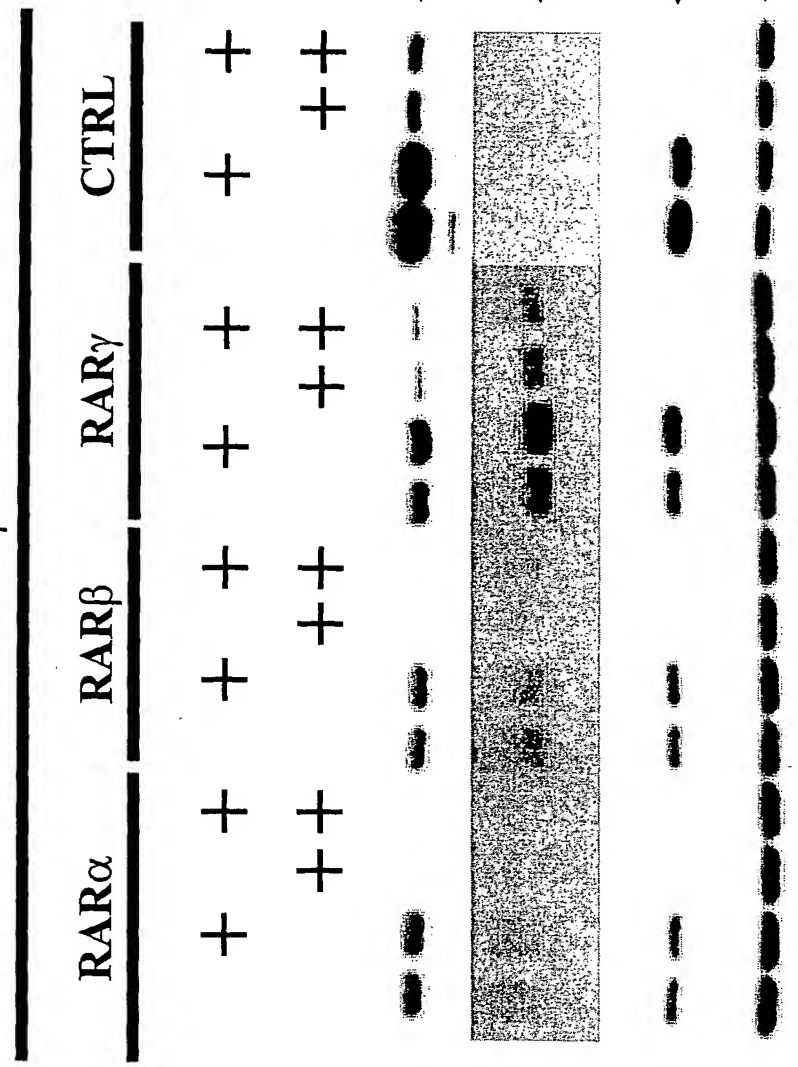


Fig. 5A

RXRα Deletion Mutants

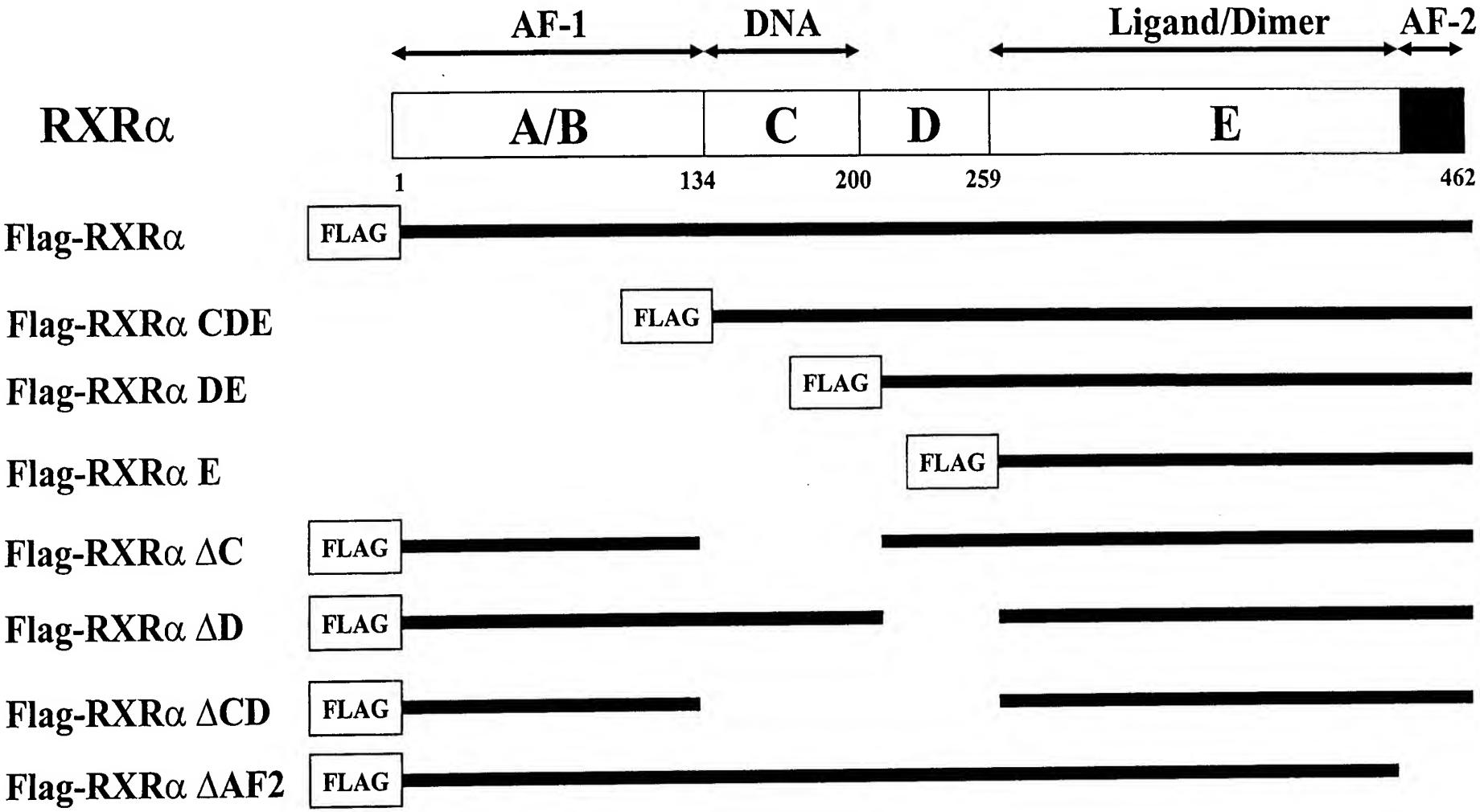


Fig. 5B

Integrity of RXRα Is Required for AGN194204-induced Reduction of RXRα & β-Catenin Proteins

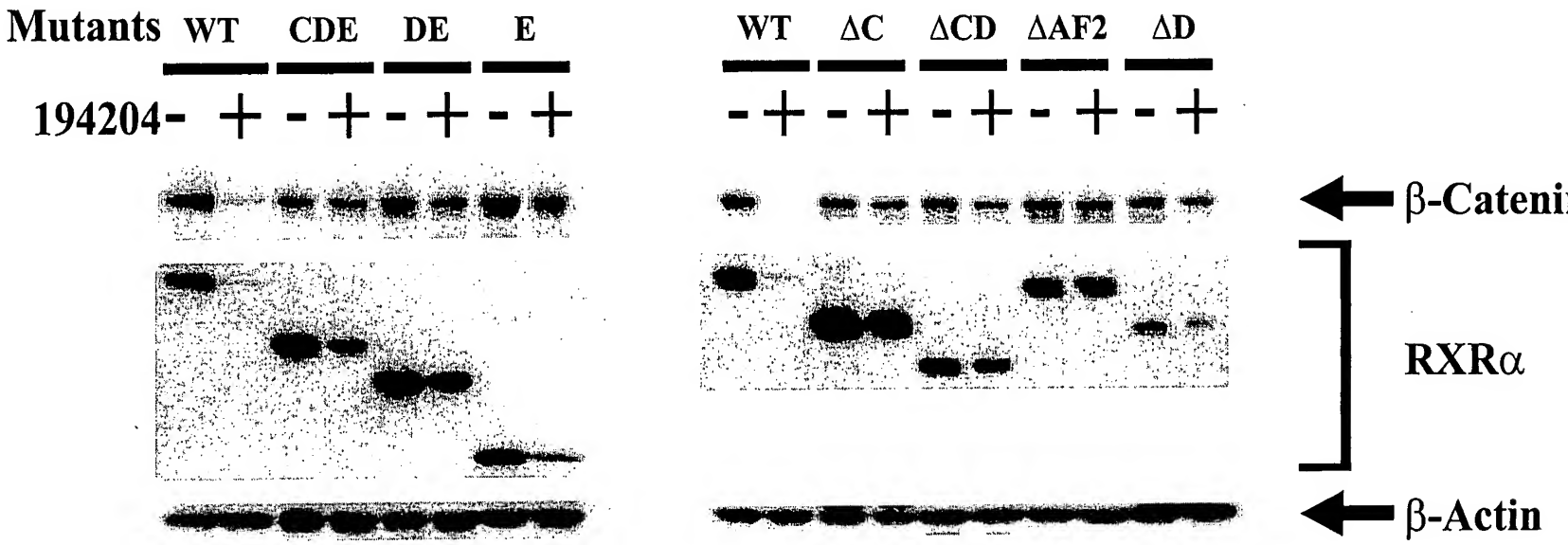


Fig. 5C

Ability of RXRα Mutants in Transactivation

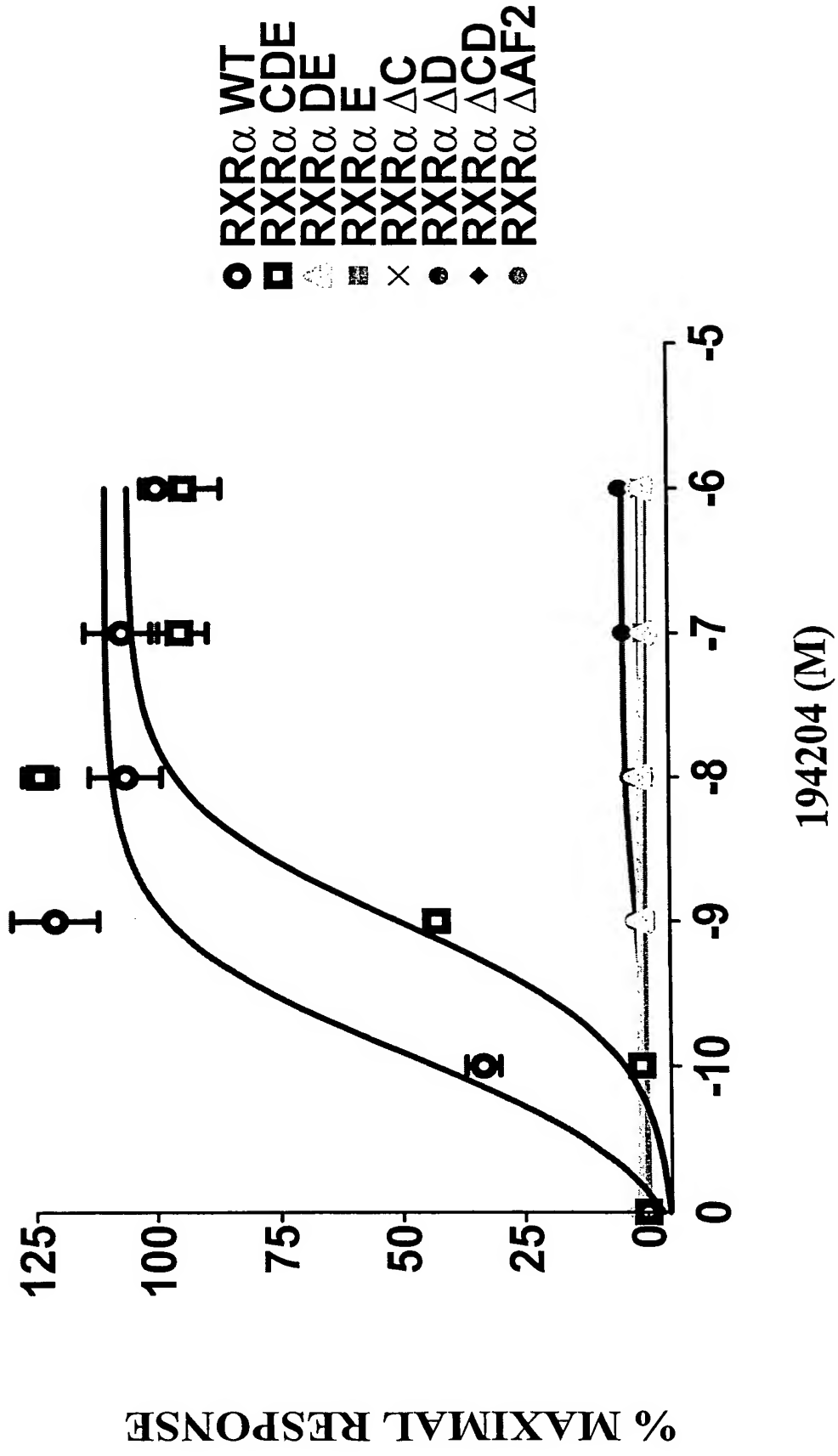
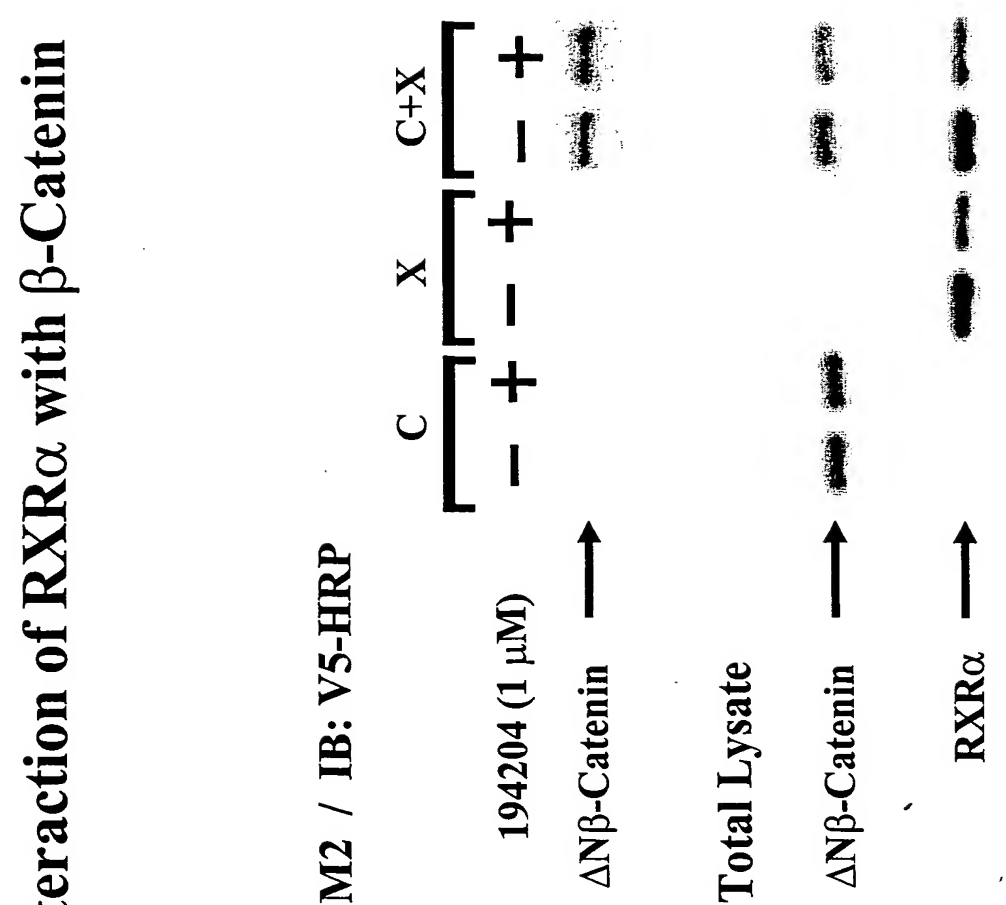


Fig. 6


Interaction of RXRα with β-Catenin
IP: M2 / IB: V5-HRP
C
X
C+X
194204 (1 μM)
– + – + – +
ΔNβ-Catenin
IB: Total Lysate
ΔNβ-Catenin
RXRα